





From the INTERNATIONAL SEARCHING AUTHORITY DUE DATE: NOTIFICATION OF TRANSMITTAL OF AMERSHAM PLC THE INTERNATIONAL SEARCH REPORT Attn. Canning, Lewis Red FORMALITIES: OR THE DECLARATION HISL Amersham Place LRCIF Little Chalfont PAT. OFF: (PCT Rule 44.1) Buckinghamshire HP7 9NA UNITED KINGDOM ON DB: P20282 CASE NO: Date of mailing (day/month/year) 11/03/2004 Applicant's or agent's file reference THROMBUS FOR FURTHER ACTION See paragraphs 1 and 4 below PZ0282-PCT International application No. International filing date (day/month/year) PCT/GB 03/04573 24/10/2003 Applicant AMERSHAM PLC 1. |X| The applicant is hereby notified that the International Search Report has been established and is transmitted herewith. Filing of amendments and statement under Article 19: The applicant is entitled, if he so wishes, to amend the claims of the International Application (see Rule 46): When? The time limit for filing such amendments is normally 2 months from the date of transmittal of the International Search Report; however, for more details, see the notes on the accompanying sheet. Where? Directly to the International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Fascimile No.: (41-22) 740.14.35 For more detailed instructions, see the notes on the accompanying sheet. The applicant is hereby notified that no International Search Report will be established and that the declaration under Article 17(2)(a) to that effect is transmitted herewith. With regard to the protest against payment of (an) additional fee(s) under Rule 40.2, the applicant is notified that: the protest together with the decision thereon has been transmitted to the International Bureau together with the applicant's request to forward the texts of both the protest and the decision thereon to the designated Offices. no decision has been made yet on the protest; the applicant will be notified as soon as a decision is made. 4. Further action(s): The applicant is reminded of the following: Shortly after 18 months from the priority date, the international application will be published by the International Bureau. If the applicant wishes to avoid or postpone publication, a notice of withdrawal of the international application, or of the priority claim, must reach the International Bureau as provided in Rules 90bis.1 and 90bis.3, respectively, before the completion of the technical preparations for international publication. Within 19 months from the priority date, a demand for international preliminary examination must be filed if the applicant wishes to postpone the entry into the national phase until 30 months from the priority date (in some Offices even later). Within 20 months from the priority date, the applicant must perform the prescribed acts for entry into the national phase before all designated Offices which have not been elected in the demand or in a later election within 19 months from the

Name and mailing address of the International Searching Authority

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priority date or could not be elected because they are not bound by Chapter II.

Natalia Morancho Alcaine

Authorized officer

These Notes are intended to give the basic instructions concerning the filing of amendments under article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article", "Rule", and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions respectively.

INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international poulication. Furthermore, it should be emphasized that provisional protection is available in some States only.

What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

When?

Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Rule 46.1).

Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been is filed, see below.

How?

Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the International application is to be published.

What documents must/may accompany the amendments?

Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- (i) the claim is unchanged;
- (ii) the claim is cancelled:
- (iii) the claim is new;
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- [Where originally there were 48 claims and after amendment of some claims there are 51]:
 Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers; claims 30, 33 and 36 unchanged; new claims 49 to 51 added.
- [Where originally there were 15 claims and after amendment of all claims there are 11]: "Claims 1 to 15 replaced by amended claims 1 to 11."
- [Where originally there were 14 claims and the amendments consist in cancelling some claims and in adding new claims]:
 "Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added." or
 "Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged."
- 4. [Where various kinds of amendments are made]: "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claim 14; claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

"Statement under article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on this description and the drawings (which cannot be amended under Article 19(1)).

The statement will be published with the international application and the amended claims.

It must be in the language in which the international appplication is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the letter indicating the differences between the claims as filed and as amended. It must be filed on a separate sheet and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

Consequence if a demand for international preliminary examination has already been filed

If, at the time of filing any amendments under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the same time of filing the amendments with the International Bureau, also file a copy of such amendments with the International Preliminary Examining Authority (see Rule 62.2(a), first sentence).

Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, where upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

For further details on the requirements of each designated/elected Office, see Volume II of the PCT Applicant's Guide.

PCT



(PCT Article 18 and Rules 43 and 44)

	A w in the couper's the reference	FOR FURTHER see Notification of Transmittal of International Search Report (Form PCT/ISA/220) as well as, where applicable, item 5 below.				
ı	troom to ecology to ation No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)			
	FT H + 04573	24/10/2003	25/10/2002			
	ACCE and AMERICAN FL.					
, ,	This triberouth row becompanied by a copy of each prior art document cited in this report.					
ı	Basis of the report					
	a - With المراجعة a - With It was filed, unless otherwise indicated under this item.					
the international search was carried out on the basis of a translation of the international application furnished to this Authority (Rule 23.1(b)).						
	b Web ಗಳ್ಳಾತರ to any nucleotide and was a arted out on the basis of the	I/or amino acid sequence disclosed in the in sequence listing ;	ternational application, the international search			
	 -,	nal application in written form.				
	=	national application in computer readable form	n.			
-		this Authority in written form.				
-		this Authority in computer readble form. sequently furnished written sequence listing de	and the displacing in the			
	international application as	filed has been furnished.				
	the statement that the infor furnished	mation recorded in computer readable form is	sidentical to the written sequence listing has been			
	2. Certain claims were foun	d unsearchable (See Box I).				
1	3. Unity of invention is lack	ing (see Box II).				
	4. With regard to the table.					
	the text is approved as submitted by the applicant.					
1		ed by this Authority to read as follows:				
	CONJUGATES OF TC COMPLI DIAGNOSTIC	EXES AND TARGETING MOIETIES	AND THEIR USE IN MRI			
	5. With regard to the abstract,					
	the text is approved as sub the text has been establish within one month from the	mitted by the applicant. ed, according to Rule 38.2(b), by this Authorit date of mailing of this international search rep	y as it appears in Box III. The applicant may, ort, submit comments to this Authority.			
	The figure of the drawings to be public					
	as suggested by the applic		X None of the figures.			
	because the applicant faile					
l	because this figure better of	characterizes the invention.				

INTERNATIONAL SEARCH REPORT

Internationa	Application No
PCT/	3/04573

A. CLASSIFICATION OF SUBJECT MATTER . IPC 7 A61K51/08

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC $\,7\,$ A61K $\,^{\circ}$

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal

C. DOCUME	C. DOCUMENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.	
X	WO 02 070018 A (MENDIZABAL MARIVI; AMERSHAM PLC (GB); DIXON MARK (GB); STOREY ANTH) 12 September 2002 (2002-09-12)	1-6, 10-13, 15-29	
Υ	See compounds at page 19-21	1-29	
	page 24, line 26 -page 27, line 7 * See in particular: example 6, example 7; page 26, lines 10-15 and example 9 "effects of temperature on labelling", page 26, line 29 * page 14, line 11-16 claims; figure 2	-	

Further documents are listed in the continuation of box C.	Patent family members are listed in annex.
Special categories of cited documents: 'A' document defining the general state of the art which is not considered to be of particular relevance 'E' earlier document but published on or after the international filing date 'L' document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) 'O' document referring to an oral disclosure, use, exhibition or other means 'P' document published prior to the international filing date but later than the priority date claimed	 *T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention *X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone *Y* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. *&* document member of the same patent family
Date of the actual completion of the international search 4 March 2004	Date of mailing of the international search report 11/03/2004

Authorized officer

Veronese, A

Form PCT/ISA/210 (second sheet) (July 1992)

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Name and mailing address of the tSA

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INTERNATIONAL SEARCH REPORT

International Application No PCT 03/04573

		PC1 03/04573		
C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT				
Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.		
X Y	WO 01 77145 A (INDREVOLL BAARD; CUTHBERTSON ALAN (NO); NYCOMED IMAGING AS (NO)) 18 October 2001 (2001-10-18) examples 2,3 page 39, line 30 -page 42, line 22 * See compound 2e, 2g page 42, line 20-22 page 39, line 20-25 claims	1-6, 10-13, 17-24 1-29		
Ÿ	WO 99 60018 A (MENDIZABAL MARIVI ;WILSON IAN ANDREW (GB); GIBSON ALEX (GB); CHAMP) 25 November 1999 (1999-11-25) page 15-17 example 4 page 22, line 19 -page 23, line 10	1-29		
Y	WO 01 89584 A (KLAVENESS JO ;NYCOMED IMAGING AS (NO); TOLLESHAUG HELGE (NO)) 29 November 2001 (2001-11-29) claims; example 2	1-6, 10-13, 17,18		
Y	WO 02 26776 A (CUTHBERTSON ALAN; NYCOMED IMAGING AS (NO)) 4 April 2002 (2002-04-04) * See compound 1 at page 27 * example 1 page 20, paragraph 2	1-6, 10-13, 17,18		

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Box I	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This Inte	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. χ	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
	Although claim 29 is directed to a diagnostic method practised on the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2.	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
; 3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Inte	ernational Searching Authority found multiple inventions in this international application, as follows:
1.	As all required additional search fees were timely paid by the applicant, this international Search Report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report
	covers only those claims for which fees were paid, specifically claims Nos.:
S	
4.	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark	on Protest The additional search fees were accompanied by the applicant's protest.
	No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

PC1 03/04573 Patent document Publication Patent family Publication cited in search report date member(s) date WO 02070018 Α 12-09-2002 CA 2439579 A1 12-09-2002 ΕP 1368064 A2 10-12-2003 WO 02070018 A2 12-09-2002 WO 0177145 Α 18-10-2001 ΑU 5068301 A 23-10-2001 CA 2405469 A1 18-10-2001 CN 1436195 T 13-08-2003 EP 1272507 A2 08-01-2003 JP 2003531835 T 28-10-2003 WO 0177145 A2 18-10-2001 US 2003204049 A1 30-10-2003 WO 9960018 Α 25-11-1999 ΑU 762736 B2 03-07-2003 AU 3942899 A 06-12-1999 BR 9910468 A 09-01-2001 CA 2332277 A1 25-11-1999 CN 1309661 T 22-08-2001 1077998 A1 EP 28-02-2001 WO 9960018 A1 25-11-1999 HU 0102154 A2 28-09-2001 JP 2002515510 T 28-05-2002 NO 20005751 A 08-01-2001 RU 2205186 C2 27-05-2003 ZA 200006313 A 17-07-2002 WO 0189584 Α 29-11-2001 ΑU 7468301 A 03-12-2001 EP 1283728 A2 19-02-2003 JP 2003534297 T 18-11-2003 WO 0189584 A2 29-11-2001 US 2003170173 A1 11-09-2003 WO 0226776 04-04-2002 Α AU 9245601 A 08-04-2002 CA 2420577 A1 04-04-2002 CN 1462276 T 17-12-2003 ΕP 1358206 A2 05-11-2003 WO 0226776 A2 04-04-2002 NO 20031323 A 24-03-2003 US 2003176639 A1 18-09-2003

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I Application No